MAY 2 8 2002 W

SEQUENCE LISTING

<110> DENEFLE, Patrice
 ROSIER-MONTUS, Marie-Francoise
 PRADES, Catherine
 ARNOULD-REGUIGNE, Isabelle
 DUVERGER, Nicolas
 ALLIKMETS, Rando
 DEAN, Michael

<120> NUCLEIC ACIDS OF THE HUMAN ABCA5, ABCA6, ABCA9, AND ABCA10 GENES, VECTORS CONTAINING SUCH NUCLEIC ACIDS, AND USES THEREOF

<130> ABCA5, 6, 9, 10

<140> US 10/005,338

<141> 2001-12-07

<150> US 60/263,231

<151> 2001-01-23

<150> FR 00403440.1

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<170> PatentIn Ver. 2.1

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Page 2

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560 545 550 555 Pro Gln Leu Asp Ile His Phe Asp Val Leu Thr Val Glu Glu Asn 565 570 575 Leu Ser Ile Leu Ala Ser Ile Lys Gly Ile Pro Ala Asn Asn Ile Ile 580 585 590 Gln Glu Val Gln Lys Val Leu Leu Asp Leu Asp Met Gln Thr Ile Lys 595 600 605 Asp Asn Gln Ala Lys Lys Leu Ser Gly Gly Gln Lys Arg Lys Leu Ser 610 620 Leu Gly Ile Ala Val Leu Gly Asn Pro Lys Ile Leu Leu Leu Asp Glu 625 630 635 640 Pro Thr Ala Gly Met Asp Pro Cys Ser Arg His Ile Val Trp Asn Leu 645 650 655 Leu Lys Tyr Arg Lys Ala Asn Arg Val Thr Val Phe Ser Thr His Phe 660 665 670 Met Asp Glu Ala Asp Ile Leu Ala Asp Arg Lys Ala Val Ile Ser Gln 675 680 685 Gly Met Leu Lys Cys Val Gly Ser Ser Met Phe Leu Lys Ser Lys Trp 690 695 700 Gly Ile Gly Tyr Arg Leu Ser Met Tyr Ile Asp Lys Tyr Cys Ala Thr 705 710 715 720 Glu Ser Leu Ser Ser Leu Val Lys Gln His Ile Pro Gly Ala Thr Leu 725 730 735 Leu Gln Gln Asn Asp Gln Gln Leu Val Tyr Ser Leu Pro Phe Lys Asp 740 745 750 Met Asp Lys Phe Ser Gly Leu Phe Ser Ala Leu Asp Ser His Ser Asn 765 760 765 Leu Gly Val Ile Ser Tyr Gly Val Ser Met Thr Thr Leu Glu Asp Val 770 780 Phe Leu Lys Leu Glu Val Glu Ala Glu Ile Asp Gln Ala Asp Tyr Ser 785 790 795 800 Val Phe Thr Gln Gln Pro Leu Glu Glu Met Asp Ser Lys Ser Phe 805 810 815 Asp Glu Met Glu Gln Ser Leu Leu Ile Leu Ser Glu Thr Lys Ala Ser 820 825 830 Leu Val Ser Thr Met Ser Leu Trp Lys Gln Gln Met Tyr Thr Ile Ala 835 840 845 Lys Phe His Phe Phe Thr Leu Lys Arg Glu Ser Lys Ser Val Arg Ser 850 855 860 Val Leu Leu Leu Leu Ile Phe Phe Thr Val Gln Ile Phe Met Phe 865 870 875 880 Leu Val His His Ser Phe Lys Asn Ala Val Val Pro Ile Lys Leu Val 885 890 895 Page 10

Pro Asp Leu Tyr Phe Leu Lys Pro Gly Asp Lys Pro His Lys Tyr Lys 900 905 910 Thr Ser Leu Leu Gln Asn Ser Ala Asp Ser Asp Ile Ser Asp Leu 915 920 925 Ile Ser Phe Phe Thr Ser Gln Asn Ile Met Val Thr Met Ile Asn Asp Ser Asp Tyr Val Ser Val Ala Pro His Ser Ala Ala Leu Asn Val Met His Ser Glu Lys Asp Tyr Val Phe Ala Ala Val Phe Asn Ser Thr Met Val Tyr Ser Leu Pro Ile Leu Val Asn Ile Ile Ser Asn Tyr Tyr Leu 980 985 990 Tyr His Leu Asn Val Thr Glu Thr Ile Gln Ile Trp Ser Thr Pro Phe 1000 Phe Gln Glu Ile Thr Asp Ile Val Phe Lys Ile Glu Leu Tyr Phe Gln 1015 1020 Ala Ala Leu Leu Gly Ile Ile Val Thr Ala Met Pro Pro Tyr Phe Ala 1030 1035 Met Glu Asn Ala Glu Asn His Lys Ile Lys Ala Tyr Thr Gln Leu Lys 1050 1045 1055 Leu Ser Gly Leu Leu Pro Ser Ala Tyr Trp Ile Gly Gln Ala Val Val 1065 Asp Ile Pro Leu Phe Phe Ile Ile Leu Ile Leu Met Leu Gly Ser Leu 1080 1075 Leu Ala Phe His Tyr Gly Leu Tyr Phe Tyr Thr Val Lys Phe Leu Ala 1095 1100 Val Val Phe Cys Leu Ile Gly Tyr Val Pro Ser Val Ile Leu Phe Thr 1110 1115 Tyr Ile Ala Ser Phe Thr Phe Lys Lys Ile Leu Asn Thr Lys Glu Phe 1130 1135 Trp Ser Phe Ile Tyr Ser Val Ala Ala Leu Xaa Cys Ile Ala Ile Thr Glu Ile Thr Phe Phe Met Gly Tyr Thr Ile Ala Thr Ile Leu His Tyr 1155 1160 1165 Ala Phe Cys Ile Ile Ile Pro Ile Tyr Pro Leu Leu Gly Cys Leu Ile 1175 1180 Ser Phe Ile Lys Ile Ser Trp Lys Asn Val Arg Lys Asn Val Asp Thr 1195 Tyr Asn Pro Trp Asp Arg Leu Ser Val Ala Val Ile Ser Pro Tyr Leu 1205 1210 1215 1210 Gln Cys Val Leu Trp Ile Phe Leu Leu Gln Tyr Tyr Glu Lys Lys Tyr

1220

1225

- Gly Gly Arg Ser Ile Arg Lys Asp Pro Phe Phe Arg Asn Leu Ser Thr 1235 1240 1245
- Lys Ser Lys Asn Arg Lys Leu Pro Glu Pro Pro Asp Asn Glu Asp Glu 1250 1255 1260
- Asp Glu Asp Val Lys Ala Glu Arg Leu Lys Val Lys Glu Leu Met Gly 1265 1270 1280
- Cys Gln Cys Cys Glu Glu Lys Pro Ser Ile Met Val Ser Asn Leu His 1285 1290 1295
- Lys Glu Tyr Asp Asp Lys Lys Asp Phe Leu Leu Ser Arg Lys Val Lys 1300 1305 1310
- Lys Val Ala Thr Lys Tyr Ile Ser Phe Cys Val Lys Lys Gly Glu Ile 1315 1320 1325
- Leu Gly Leu Leu Gly Pro Asn Gly Ala Gly Lys Ser Thr Ile Ile Asn 1330 1340
- Ile Leu Val Gly Asp Ile Glu Pro Thr Ser Gly Gln Val Phe Leu Gly 1345 1350 1360
- Asp Tyr Ser Ser Glu Thr Ser Glu Asp Asp Ser Leu Lys Cys Met 1365 1370 1375
- Gly Tyr Cys Pro Gln Ile Asn Pro Leu Trp Pro Asp Thr Thr Leu Gln 1380 1385 1390
- Glu His Phe Glu Ile Tyr Gly Ala Val Lys Gly Met Ser Ala Ser Asp 1395 1400
- Met Lys Glu Val Ile Ser Arg Ile Thr His Ala Leu Asp Leu Lys Glu 1410 1420
- His Leu Gln Lys Thr Val Lys Lys Leu Pro Ala Gly Ile Lys Arg Lys 1425 1430 1435 1440
- Leu Cys Phe Ala Leu Ser Met Leu Gly Asn Pro Gln Ile Thr Leu Leu 1445 1450 1455
- Asp Glu Pro Ser Thr Gly Met Asp Pro Lys Ala Lys Gln His Met Trp 1460 1465 1470
- Arg Ala Ile Arg Thr Ala Phe Lys Asn Arg Lys Arg Ala Ala Ile Leu 1475 1480 1485
- Thr Thr His Tyr Met Glu Glu Ala Glu Ala Val Cys Asp Arg Val Ala 1490 1495 1500
- Ile Met Val Ser Gly Gln Leu Arg Cys Ile Gly Thr Val Gln His Leu 1505 1510 1515 1520
- Lys Ser Lys Phe Gly Lys Gly Tyr Phe Leu Glu Ile Lys Leu Lys Asp 1525 1530 1535
- Trp Ile Glu Asn Leu Glu Val Asp Arg Leu Gln Arg Glu Ile Gln Tyr 1540 1545 1550
- Ile Phe Pro Asn Ala Ser Arg Gln Glu Ser Phe Ser Ser Ile Leu Ala 1555 1560 1565
- Tyr Lys Ile Pro Lys Glu Asp Val Gln Ser Leu Ser Gln Ser Phe Phe Page 12

Lys Leu Glu Glu Ala Lys His Ala Phe Ala Ile Glu Glu Tyr Ser Phe 1590 Ser Gln Ala Thr Leu Glu Gln Val Phe Val Glu Leu Thr Lys Glu Gln 1610 Glu Glu Glu Asp Asn Ser Cys Gly Thr Leu Asn Ser Thr Leu Trp Trp Glu Arg Thr Gln Glu Asp Arg Val Val Phe 1640

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His Pro Val Met Glu Glu Leu Met Ser Val Thr Ala Ile Thr Met Lys 195

Thr Leu Pro Phe Ile Thr Lys Asn Leu Leu His Asn Glu Met Phe Ile 210 215 220

Leu Phe Phe Leu Leu His Phe Ser Pro Leu Val Tyr Phe Ile Ser Leu Asn Val Thr Lys Glu Arg Lys Lys Ser Lys Asn Leu Met Lys Met Met 245 250 255 Gly Leu Gln Asp Ser Ala Phe Trp Leu Ser Trp Gly Leu Ile Tyr Ala 260 265 270 Gly Phe Ile Phe Ile Ile Ser Ile Phe Ile Thr Ile Ile Ile Thr Phe $275 \hspace{1.5cm} 280 \hspace{1.5cm} 285$ Thr Gln Ile Ile Val Met Thr Gly Phe Met Val Ile Phe Ile Leu Phe 290 295 300 Phe Leu Tyr Gly Leu Ser Leu Val Ala Leu Val Phe Leu Met Ser Val 305 310 315 320 305 Leu Leu Lys Lys Ala Val Leu Thr Asn Leu Val Val Phe Leu Leu Thr 325 330 335 Leu Phe Trp Gly Cys Leu Gly Phe Thr Val Phe Tyr Glu Gln Leu Pro 340 345 350 Ser Ser Leu Glu Trp Ile Leu Asn Ile Cys Ser Pro Phe Ala Phe Thr 355 360 365 Thr Gly Met Ile Gln Ile Ile Lys Leu Asp Tyr Asn Leu Asn Gly Val 370 375 380 Ile Phe Pro Asp Pro Ser Gly Asp Ser Tyr Thr Met Ile Ala Thr Phe 385 390 395 400 Ser Met Leu Leu Asp Gly Leu Ile Tyr Leu Leu Ala Leu Tyr 405 410 415 Phe Asp Lys Ile Leu Pro Tyr Gly Asp Glu Arg His Tyr Ser Pro Leu 420 425 430 Phe Leu Asn Ser Ser Ser Cys Phe Gln His Gln Arg Thr Asn Ala 435 440 445 Lys Val Ile Glu Lys Glu Ile Asp Ala Glu His Pro Ser Asp Asp Tyr 450 455 460 Phe Glu Pro Val Ala Pro Glu Phe Gln Gly Lys Glu Ala Ile Arg Ile 465 470 475 480 Arg Asn Val Lys Lys Glu Tyr Lys Gly Lys Ser Gly Lys Val Glu Ala 485 490 495 Leu Lys Gly Leu Leu Phe Asp Ile Tyr Glu Gly Gln Ile Thr Ala Ile 500 510 Leu Gly His Ser Gly Ala Gly Lys Ser Ser Leu Leu Asn Ile Leu Asn 515 520 525 Gly Leu Ser Val Pro Thr Glu Gly Ser Val Thr Ile Tyr Asn Lys Asn 530 540 Leu Ser Glu Met Gln Asp Leu Glu Glu Ile Arg Lys Ile Thr Gly Val 545 550 555 560 Cys Pro Gln Phe Asn Val Gln Phe Asp Ile Leu Thr Val Lys Glu Asn Page 14

Leu Ser Leu Phe Ala Lys Ile Lys Gly Ile His Leu Lys Glu Val Glu 580 585 590 Gln Glu Val Gln Arg Ile Leu Leu Glu Leu Asp Met Gln Asn Ile Gln 600 Asp Asn Leu Ala Lys His Leu Ser Glu Gly Gln Lys Arg Lys Leu Thr 610 615 620 Phe Gly Ile Thr Ile Leu Gly Asp Pro Gln Ile Leu Leu Leu Asp Glu 625 630 635 640 Pro Thr Thr Gly Leu Asp Pro Phe Ser Arg Asp Gln Val Trp Ser Leu 645 650 655 Leu Arg Glu Arg Arg Ala Asp His Val Ile Leu Phe Ser Thr Gln Ser 660 670 Met Asp Glu Ala Asp Ile Leu Ala Asp Arg Lys Val Ile Met Ser Asn 675 680 685 Gly Arg Leu Lys Cys Ala Gly Ser Ser Met Phe Leu Lys Arg Arg Trp 690 695 700 Gly Leu Gly Tyr His Leu Ser Leu His Arg Asn Glu Ile Cys Asn Pro 705 710 715 720 Glu Gln Ile Thr Ser Phe Ile Thr His His Ile Pro Asp Ala Lys Leu 725 730 735 Lys Thr Glu Asn Lys Glu Lys Leu Val Tyr Thr Leu Pro Leu Glu Arg 740 745 750 Thr Asn Thr Phe Pro Asp Leu Phe Ser Asp Leu Asp Lys Cys Ser Asp 765 Gln Gly Val Thr Gly Tyr Asp Ile Ser Met Ser Thr Leu Asn Glu Val 770 775 780 Phe Met Lys Leu Glu Gly Gln Ser Thr Ile Glu Gln Asp Phe Glu Gln 785 790 795 800 Val Glu Met Ile Arg Asp Ser Glu Ser Leu Asn Glu Met Glu Leu Ala 805 810 815 His Ser Ser Phe Ser Glu Met Gln Thr Ala Val Ser Asp Met Gly Leu 825 Trp Arg Met Gln Val Phe Ala Met Ala Arg Leu Arg Phe Leu Lys Leu 835 840 845 Lys Arg Gln Thr Lys Val Leu Leu Thr Leu Leu Leu Val Phe Gly Ile 850 855 860 Ala Ile Phe Pro Leu Ile Val Glu Asn Ile Ile Tyr Ala Met Leu Asn 865 870 875 880 Glu Lys Ile Asp Trp Glu Phe Lys Asn Glu Leu Tyr Phe Leu Ser Pro 885 890 895 Gly Gln Leu Pro Gln Glu Pro Arg Thr Ser Leu Leu Ile Ile Asn Asn 905 910 Page 15

Thr	Glu	Ser 915	Asn	Ile	Glu	Asp	Phe 920	Ile	Lys	Ser	Leu	Lys 925	His	Gln	Asn
Ile	Leu 930	Leu	Glu	٧al	Asp	Asp 935	Phe	Glu	Asn	Arg	Asn 940	Gly	Thr	Asp	Gly
Leu 945	Ser	Tyr	Asn	Gly	Ala 950	Ile	Ile	٧a٦	Ser	Gly 955	Lys	Gln	Lys	Asp	Tyr 960
Arg	Phe	Ser	٧a٦	va1 965	Cys	Asn	Thr	Lys	Arg 970	Leu	His	Cys	Phe	Pro 975	Ile
Leu	Met	Asn	11e 980	Ile	Ser	Asn	Gly	Leu 985	Leu	Gln	Met	Phe	Asn 990	His	Thr
Gln	His	11e 995	Arg	Ile	Glu		Ser 1000	Pro	Phe	Pro		Ser 1005	His	Ile	Gly
	Trp 1010	Thr	Gly	Leu		Asp 1015	Gly	Ser	Phe		Leu 1020	Phe	Leu	val	Leu
Cys 102		Ile	Ser		Tyr 1030	Ile	Thr	Met		Ser 1035	Ile	Ser	Asp	Tyr	Lys 1040
Lys	Asn	Ala	Lys	Ser 1045	Gln	Leu	Trp		Ser 1050	Gly	Leu	Tyr		Ser 1055	Ala
Tyr	Trp		Gly 1060	Gln	Ala	Leu		Asp 1065	val	Ser	Phe	Phe	Ile 1070	Leu	Ile
Leu		Leu 1075	Met	Tyr	Leu		Phe 1080	Tyr	Ile	Glu		Меt 1085	Gln	Tyr	Leu
	Ile 1090	Thr	Ser	Gln		val 1095	Phe	Ala	Leu		Ile 1100	٧a٦	Thr	Pro	Gly
Tyr 110		Ala	Ser		Val 1110	Phe	Phe	Ile		Меt 1115	Ile	Ser	Phe		Phe 1120
Arg	Lys	Arg	Arg	Lys 1125	Asn	Ser	Gly		Trp 1130	Ser	Phe	Tyr		Phe 1135	Phe
Ala	Ser		Ile 1140	Met	Phe	Ser		Thr 1145	Leu	Ile	Asn		Phe 1150	Asp	Leu
Ser		Leu 1155	Ile	Thr	Thr		∨a1 1160	Leu	٧a٦	Pro		Туг 1165	Thr	Leu	Leu
	Phe L170	Lys	Thr	Phe		Glu 1175	val	Arg	Asp		Glu 1180	His	Tyr	Arg	Glu
Phe 1185		Glu	Ala		Phe 1190	Glu	Leu	Ser		Thr 1195	Asp	Phe	Leu	۷al	Cys 1200
Phe	Ile	Pro	Tyr	Phe 1205	Gln	Thr	Leu		Phe 1210	٧a٦	Phe	٧a٦		Arg 1215	Cys
Met	Glu		Lys 1220	Cys	Gly	Lys		Arg 1225	Met	Arg	Lys		Pro 1230	٧al	Phe
Arg		Ser 1235	Pro	Gln	Ser		Asp 1240	Аlа	Lys	Pro		Pro 1245	Glu	Glu	Pro

- Ala Leu Thr Thr Ser Ile Leu Asp Glu Lys Pro Val Ile Ile Ala Ser 1265 1270 1275 1280
- Cys Leu His Lys Glu Tyr Ala Gly Gln Lys Lys Ser Cys Phe Ser Lys 1285 1290 1295
- Arg Lys Lys Ile Ala Ala Arg Asn Ile Ser Phe Cys Val Gln Glu 1300 1305 1310
- Gly Glu Ile Leu Gly Leu Leu Gly Pro Ser Gly Ala Gly Lys Ser Ser 1315 1320 1325
- Ser Ile Arg Met Ile Ser Gly Ile Thr Lys Pro Thr Ala Gly Glu Val 1330 1340
- Glu Leu Lys Gly Cys Ser Ser Val Leu Gly His Leu Gly Tyr Cys Pro 1345 1350 1355 1360
- Gln Glu Asn Val Leu Trp Pro Met Leu Thr Leu Arg Glu His Leu Glu 1365 1370 1375
- Val Tyr Ala Ala Val Lys Gly Leu Arg Lys Ala Asp Ala Arg Leu Ala 1380 1385 1390
- Ile Ala Arg Leu Val Ser Ala Phe Lys Leu His Glu Gln Leu Asn Val 1395 1400 1405
- Pro Val Gln Lys Leu Thr Ala Gly Ile Thr Arg Lys Leu Cys Phe Val 1410 1415 1420
- Leu Ser Leu Leu Gly Asn Ser Pro Val Leu Leu Leu Asp Glu Pro Ser 1425 1430 1435 1440
- Thr Gly Ile Asp Pro Thr Gly Gln Gln Met Trp Gln Ala Ile Gln
 1445 1450 1455
- Ala Val Val Lys Asn Thr Glu Arg Gly Val Leu Leu Thr Thr His Asn 1460 1465 1470
- Leu Ala Glu Ala Glu Ala Leu Cys Asp Arg Val Ala Ile Met Val Ser 1475 1480 1485
- Gly Arg Leu Arg Cys Ile Gly Ser Ile Gln His Leu Lys Asn Lys Leu 1490 1495 1500
- Gly Lys Asp Tyr Ile Leu Glu Leu Lys Val Lys Glu Thr Ser Gln Val 1505 1510 1515 1520
- Thr Leu Val His Thr Glu Ile Leu Lys Leu Phe Pro Gln Ala Ala Gly 1525 1530 1535
- Gln Glu Arg Tyr Ser Ser Leu Leu Thr Tyr Lys Leu Pro Val Ala Asp 1540 1545 1550
- Val Tyr Pro Leu Ser Gln Thr Phe His Lys Leu Glu Ala Val Lys His 1555 1560 1565
- Asn Phe Asn Leu Glu Glu Tyr Ser Leu Ser Gln Cys Thr Leu Glu Lys 1570 1575 1580
- Val Phe Leu Glu Leu Ser Lys Glu Gln Glu Val Gly Asn Phe Asp Glu Page 17

1585 1590 1595

Glu Ile Asp Thr Thr Met Arg Trp Lys Leu Leu Pro His Ser Asp Glu 1605 1610 1615

1600

Pro

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Glu Trp Leu Phe Ser Phe Leu Leu Val Leu Phe Leu Tyr Leu Phe Phe 35 40 45

Ser Asn Leu His Gln Val His Asp Thr Pro Gln Met Ser Ser Met Asp 50 55 60

Leu Gly Arg Val Asp Ser Phe Asn Asp Thr Asn Tyr Val Ile Ala Phe 65 70 75 80

Ala Pro Glu Ser Lys Thr Thr Gln Glu Ile Met Asn Lys Val Ala Ser 85 90 95

Ala Pro Phe Leu Lys Gly Arg Thr Ile Met Gly Trp Pro Asp Glu Lys $100 \hspace{1cm} 105 \hspace{1cm} 110$

Ser Met Asp Glu Leu Asp Leu Asn Tyr Ser Ile Asp Ala Val Arg Val 115 120 125

Ile Phe Thr Asp Thr Phe Ser Tyr His Leu Lys Phe Ser Trp Gly His 130 135 140

Arg Ile Pro Met Met Lys Glu His Arg Asp His Ser Ala His Cys Gln 145 150 155 160

Ala Val Asn Glu Lys Met Lys Cys Glu Gly Ser Glu Phe Trp Glu Lys 165 170 175

Gly Phe Val Ala Phe Gln Ala Ala Ile Asn Ala Ala Ile Ile Glu Ile 180 185 190

Ala Thr Asn His Ser Val Met Glu Gln Leu Met Ser Val Thr Gly Val 195 200 205

His Met Lys Ile Leu Pro Phe Val Ala Gln Gly Gly Val Ala Thr Asp 210 215 220

Phe Phe Ile Phe Phe Cys Ile Ile Ser Phe Ser Thr Phe Ile Tyr Tyr 225 230 235 240

Val Ser Val Asn Val Thr Gln Glu Arg Gln Tyr Ile Thr Ser Leu Met 245 250 255

Thr Met Met Gly Leu Arg Glu Ser Ala Phe Trp Leu Ser Trp Gly Leu 265 Met Tyr Ala Gly Phe Ile Leu Ile Met Ala Thr Leu Met Ala Leu Ile 275 280 285 Lys Ser Ala Gln Ile Val Val Leu Thr Gly Phe Val Met Val Phe 290 295 300 Thr Leu Phe Leu Leu Tyr Gly Leu Ser Leu Ile Thr Leu Ala Phe Leu 305 310 315 320 Met Ser Val Leu Ile Lys Lys Pro Phe Leu Thr Gly Leu Val Val Phe 325 330 335 Leu Leu Ile Val Phe Trp Gly Ile Leu Gly Phe Pro Ala Leu Tyr Thr 345 His Leu Pro Ala Phe Leu Glu Trp Thr Leu Cys Leu Leu Ser Pro Phe Phe Thr Val Gly Met Ala Gln Leu Ile His Leu Asp Tyr Asp Val 370 375 380 Asn Ser Asn Ala His Leu Asp Ser Ser Gln Asn Pro Tyr Leu Ile Ile Ala Thr Leu Phe Met Leu Val Phe Asp Thr Leu Leu Tyr Leu Val Leu Thr Leu Tyr Phe Asp Lys Ile Leu Pro Ala Glu Tyr Gly His Arg Cys Pro Leu Phe Phe Leu Lys Ser Cys Phe Trp Phe Gln His Gly Arg Ala Asn His Val Val Leu Glu Asn Glu Thr Asp Ser Asp Pro Thr Pro Asn Asp Cys Phe Glu Pro Val Ser Pro Glu Phe Cys Gly Lys Glu Ala Ile Arg Ile Lys Asn Leu Lys Lys Glu Tyr Ala Gly Lys Cys Glu Arg 485 490 495 Val Glu Ala Leu Lys Gly Val Val Phe Asp Ile Tyr Glu Gly Gln Ile 500 505 510 Thr Ala Leu Leu Gly His Ser Gly Ala Gly Lys Thr Thr Leu Leu Asn 515 520 525 Ile Leu Ser Gly Leu Ser Val Pro Thr Ser Gly Ser Val Thr Val Tyr 530 540 Asn His Thr Leu Ser Arg Met Ala Asp Ile Glu Asn Ile Ser Lys Phe 545 550 555 560 Thr Gly Phe Cys Pro Gln Ser Asn Val Gln Phe Gly Phe Leu Thr Val Lys Glu Asn Leu Arg Leu Phe Ala Lys Ile Lys Gly Ile Leu Pro His 580 585 590 Glu Val Glu Lys Glu Val Gln Arg Val Val Gln Glu Leu Glu Met Glu Page 19

Ile Gln Asp Ile Leu Ala Gln Asn Leu Ser Gly Gly Gln Asn Arg 610 615 620 Lys Leu Thr Phe Gly Ile Ala Ile Leu Gly Asp Pro Gln Val Leu Leu 625 630 635 640 Leu Asp Glu Pro Thr Ala Gly Leu Asp Pro Leu Ser Arg His Arg Ile 645 650 655 Trp Asn Leu Leu Lys Glu Gly Lys Ser Asp Arg Val Ile Leu Phe Ser 660 665 670 Thr Gln Phe Ile Asp Glu Ala Asp Ile Leu Ala Asp Arg Lys Val Phe 675 680 685 Ser Asn Gly Lys Leu Lys Cys Ala Gly Ser Ser Leu Phe Leu Lys 690 695 700 Lys Trp Gly Ile Gly Tyr His Leu Ser Leu His Leu Asn Glu Arg
710 715 720 Cys Asp Pro Glu Ser Ile Thr Ser Leu Val Lys Gln His Ile Ser Asp 725 730 735 Ala Lys Leu Thr Ala Gln Ser Glu Glu Lys Leu Val Tyr Ile Leu Pro 740 745 750 Leu Glu Arg Thr Asn Lys Phe Pro Glu Leu Tyr Arg Asp Leu Asp Arg 755 760 765 Cys Ser Asn Gln Gly Ile Glu Asp Tyr Gly Val Ser Ile Thr Thr Leu 770 780 Asn Glu Val Phe Leu Lys Leu Glu Gly Lys Ser Thr Ile Asp Glu Ser 785 790 795 800 Asp Ile Gly Ile Trp Gly Gln Leu Gln Thr Asp Gly Ala Lys Asp Ile 805 810 815 Gly Ser Leu Val Glu Leu Glu Gln Val Leu Ser Ser Phe His Glu Thr 820 825 830 Arg Lys Thr Ile Ser Gly Val Ala Leu Trp Arg Gln Gln Val Cys Ala 835 840 845 Ile Ala Lys Val Arg Phe Leu Lys Leu Lys Lys Glu Arg Lys Ser Leu 850 860 Trp Thr Ile Leu Leu Phe Gly Ile Ser Phe Ile Pro Gln Leu Leu 865 870 875 880 Glu His Leu Phe Tyr Glu Ser Tyr Gln Lys Ser Tyr Pro Trp Glu Leu 885 890 895 Ser Pro Asn Thr Tyr Phe Leu Ser Pro Gly Gln Gln Pro Gln Asp Pro 900 905 910 Leu Thr His Leu Leu Val Ile Asn Lys Thr Gly Ser Thr Ile Asp Asn 915 920 925 Phe Leu His Ser Leu Arg Arg Gln Asn Ile Ala Ile Glu Val Asp Ala 930 935 940 Page 20

945	Gly	Thr	Arg	Asn	Gly 950	Thr	Asp	Asp	Pro	ser 955	Tyr	Asn	Gly	Ala	960
Ile	val	Ser	Gly	Asp 965	Glu	Lys	Asp	His	Arg 970	Phe	Ser	Ile	Ala	Cys 975	Asn
Thr	Lys	Arg	Leu 980	Asn	Cys	Phe	Pro	Val 985	Leu	Leu	Asp	۷a٦	11e 990	Ser	Asn
Gly	Leu	Leu 995	Gly	Ile	Phe		Ser 1000	Ser	Glu	His		G]n 1005	Thr	Asp	Arg
	Thr L010	Phe	Phe	Glu		His 1015	Met	Asp	Tyr	Glu	Tyr 1020	Gly	Tyr	Arg	Ser
Asn 1025		Phe	Phe	Trp	11e 1030	Pro	Met	Ala		Ser 1035	Phe	Thr	Pro		11e 1040
Ala	Met	Ser		11e 1045	Gly	Asp	Tyr		Lys 1050	Lys	Аlа	His		G]n L055	Leu
Arg	Ile		G]y 1060	Leu	туr	Pro		а1а 1065	Tyr	Trp	Phe		Gln 1070	Ala	Leu
val		va1 1075	Ser	Leu	Tyr		Leu 1080	Ile	Leu	Leu		меt 1085	Gln	Ile	Met
	Tyr L090	Ile	Phe	Ser		G]u 1095	Glu	Ile	Ile		11e 1100	Ile	Gln	Asn	Leu
Leu 110	Ile 5	Gln	Ile	Leu	Cys 1110	Ser	Ile	Gly	Tyr	∨a1 1115	Ser	Ser	Leu	Val	Phe L120
Leu	Thr	Tyr		Ile 1125	Ser	Phe	Ile		Arg 1130	Asn	Gly	Arg		Asn L135	Ser
Gly	Ile		Ser 1140	Phe	Phe	Phe		Ile 1145	Val	val	Ile		Ser 1150	Ile	val
Ala		Asp 1155	Leu	Asn	Glu		Gly 1160	Phe	Leu	Gly		Phe 1165	Phe	Gly	Thr
	Leu 1170	Ile	Pro	Pro		Thr 1175	Leu	Ile	Gly		Leu 1180	Phe	Ile	Phe	Ser
Glu 1185	Ile 5	Ser	Pro	Asp	Ser 1190	Met	Asp	Tyr		Gly 1195	Ala	Ser	Glu	Ser	G1u L200
Ile	val	Tyr		А1а 1205	Leu	Leu	Ile		Туг 1210	Leu	His	Phe		11e 1215	Phe
Leu	Phe		Leu 1220	Arg	Cys	Leu		меt 1225	Asn	Cys	Arg		Lys 1230	Leu	Met
Arg		Asp L235	Pro	٧al	Phe		11e 1240	Ser	Pro	Arg		Asn 1245	Ala	Ile	Phe
	Asn L250	Pro	Glu	Glu		Glu 1255	Gly	Glu	Glu		Asp 1260	Ile	Gln	Met	Glu
Arg 1265		Arg	Thr	٧al	Asn 1270	Ala	Met	Ala		Arg 1275	Asp	Phe	Asp		Thr L280

Pro Val Ile Ile Ala Ser Cys Leu Arg Lys Glu Tyr Ala Gly Lys Lys 1285 1290 1295

Lys Asn Cys Phe Ser Lys Arg Lys Lys Thr Ile Ala Thr Arg Asn Val 1300 1305 1310

Ser Phe Cys Val Lys Lys Gly Glu Val Ile Gly Leu Leu Gly His Asn 1315 1320 1325

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Pro Thr Ala Gly Gln Val Ile Leu Lys Gly Ser Gly Gly Gly Glu Pro 1345 1350 1355 1360

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Arg Lys Gly Asp Ala Met Ile Ala Ile Thr Arg Leu Val Asp Ala Leu 1395 1400 1405

Lys Leu Gln Asp Gln Leu Lys Ala Pro Val Lys Thr Leu Ser Glu Gly 1410 1420

Ile Lys Arg Lys Leu Arg Phe Val Leu Ser Ile Leu Gly Asn Pro Ser 1425 1430 1445

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1445 1450 1455

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Gly Ala Leu Leu Thr Thr His Tyr Met Ala Glu Ala Glu Ala Val Cys 1475 1480 1485

Asp Arg Val Ala Ile Met Val Ser Gly Arg Leu Arg Cys Ile Gly Ser 1490 1495 1500

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Phe Lys Leu Glu Ile Val Lys Gln Ser Phe Asp Leu Glu Glu Tyr Ser 1570 1575 1580

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950	255	960
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